United States
Environm ental Protection
Agency

Office of Prevention, Pesticides and Toxic Substances
(7501C)



Pesticide FactSheet

Name of Chemical: A cequinocyl

Reason for Issuance: Conditional Registration

Date Issued: Septem ber 26, 2003

1. <u>DESCRIPTION OF CHEM ICAL</u>

Generic Name: 3-dodecyl-1, 4-dihydro-1, 4-dioxo-2-naphthyl-acetate

CommonName: Acequinocyl

TradeName: Kanemite

EPA PC Code: 006329

Chemical Abstracts

Service (CAS) Number: 57960-19-7

Yearof Initial

Registration: 2003

Pesticide Type: Miticide

Chemical Family: Quinoline

Function, Mode of Action The majorm etabolite of acequinocyl (deacety lated

acequinocyl) inhibits electron transfer by binding the Q o Center

at complex III in the mitochondria.

Classification of

End-Use Product This product is not a restricted use pesticide

U.S. Producer: A rvesta Corporation

2. <u>USE PATTERNS AND FORM ULATIONS</u>

Application Sites: A cequinocyl is registered for use on ornam ental plants grown in

com m ercial greenhouses and shadehouses for control of

various mites.

Types of Formulations: 96.8% technical product

15.8% soluble concentrate end-use product

Use Summary: The formulated product, which is a soluble concentrate product

containing 15.8% acequinocylis diluted in water and applied as a full coverage foliar spray "to drip" at 0.06 lb.a.i./100 gallons on roses and

impatiens and from 0.06-0.125 lb.a.i./100 gallons on other

ornam entals. The maximum application rate percrop cycle is 0.3 lb. ai/A on roses and impatiens and 0.6 lb.ai/A on other ornam entals. Successive applications of the pesticide are not recommended in order

to reduce any potential for developm ent of resistence.

3. SCIENCE FINDINGS

A cequinocyl is a mem berof the quinoline class of insecticides/miticides and has been designated as a "reduced risk" pesticide by EPA. The available product chemistry, toxicology, ecological effects and environmental fate data for acequinocyl are summarized below, along with the estimated risks to human health and the environment from its use on ornamental crops grown in commercial greenhouses and shadehouses:

Chemical Characteristics

PRO PERTY	TECHNICAL	END-USE
Color	LightBrown (TGAI) SoftYellow (PAI)	Pale Y ellow
Physical State	Flakes (TGAI) Crystals (PAI)	liquid suspension
0 dor	faintly earthy (FGAI) non-detectable (PAI)	detergent-like
0 xidation/reduction	none	no reaction
Flamm ability	nothighly flammable	no flash point
Explodability	notexplosive	notexplosive
Storage Stability	data gap	stable for 2 years

PRO PERTY	TECHNICAL	END-USE
M iscibility	NA	NA
Conosion Characteristics	data gap	non-conosive
pН	694 (TGAI)	710
V iscosity	NA	405.95 cS (20°C) 217.23 cS (40°C
Melting Point-TGAI	59.6℃	NA
Boiling Point-TGA I	could notbe m easured, TS changed over 200°C	NA
Relative density-TGAI	1.13 at20℃	1.04
D issociation constants in water-TGAI	could not be measured, low solubility in water	NA
Partition coefficient(n- octanol/water)-TGAI	log Pow 62	NA
W atersolubility; column elution m ethod; shake flask m ethod-TGAI	inwater=6.69ug/Lat20℃	NA
Vaporpressure-TGAI	1.69x10-6 Pa.at25℃	NA

Toxicology Characteristics

A . A cute Toxicity D ata on Technical A cequinocyland End-Use Product (15.8% SC)

	TECHNICAL PRODUCT		END-USE PRODUCT	
STUDY	RESULTS	TOXICITY CATEGORY	RESULTS	TOXICITY CATEGORY
acute oral toxicity; mouse; LD 50	5000 m g/kg bw	IV	5000 m g/kg bw	IV
acute oral toxicity; rat; LD 50	5000 m g/kg bw	IV	5000 m g/kg bw	IV
acute derm altoxicity; LD 50	2000 m g/kg bw	ш	5000 m g/kg bw	IV
acute inhalation; LC50	0.84 m g/L	ш	4.49 m g/L	IV
primary eye irritation	notan eye imilant	IV	no imitation at 24 hrs. or subsequently	IV

	TECHNICAL PRODUCT		END-USE PRODUCT	
STUDY	RESULTS	TOXICITY CATEGORY	RESULTS	TOXICITY CATEGORY
prim ary dem al inditation	nota dem al imitant	IV	no derm al reaction	IV
derm al sensitization	nota demal sensitizer	negative	data gap	data gap

B. Subchronic Toxicity Data

The subchronic studies (28-day derm alstudy and subchronic feeding study) used in the risk assessment for acequinocylare sum marized in the table under Toxicological Endpoints, below.

C.M utagenicity

A cequinocylwas not mutagenic in 1) Salmonella typhimurium/Escherichia coliassay, 2) in vitro mammalian cellgene mutation assay, 3) in vitro mammalian cytogenetics, and 4) a micronucleus assay with mouse erythrocytes. O verall, the data suggest that acequinocyl is negative form utagenicity in vitro and in vivo.

D. Carcinogenicity

Carcinogenicity data are required for food use pesticides and for non-food use pesticides when chronic exposure of handlers and/or by standers may occur as a result of the pesticide's use. The use of acequinocylon greenhouse/shadehouse grown or namentals is expected to result in short-and intermediate-term handler exposure but is not expected to result in chronic exposure to the pesticide. Therefore, cancer risk is not a concern for this use of acequinocyl, and carcinogenicity studies are not required.

<u>ToxicologicalEndpoints</u>

The toxicological endpoints used in the hum an health risk assessment for acequinocylare sum marized in the table below. The identified short-term (1-30 days) dermal toxicological endpoint of 200 mg a.i./kg bw /day is based upon increased blood clotting times seen in a 28-day ratdermal study. The intermediate-term (1-6 m onths) dermal endpoint is the same and is identified from the same study. A short-term (1-30 days) inhalation endpoint (30 mg a.i./kg bw /day) was identified from a subchronic ratstudy where the effect seen was reddish urine, which is indicative of the clotting abnormalities seen in the 28-day dermal study. An intermediate-term (1-6 m onths) inhalation endpoint was identified and is the same as the short-term inhalation endpoint.

Exposure Scenario	Dose Used in Risk Assessment	LevelofConcem forRisk Assessment	Study and Toxicological E ffects
Short-Term Dermal (1 to 30 days)	DemalStudy NOAEL = 200 mg/kg/day	O ccupationalLOC forMOE = 100	28-day Demal Toxicity Study in the Rat Increased clotting factor times.
Intermediate- Term Dermal (1 to 6 months)	DemalStudy NOAEL = 200 mg/kg/day	OccupationalLOC forMOE = 100	28-day Dermal Toxicity Study in the Rat Increased clotting factor times.
Short-Term Inhalation (1 to 30 days)	Oral study NOAEL=30 mg/kg/day (inhalation absorption rate= 100%)	OccupationalLOC forMOE = 100	Subchronic feeding study in the rat Reddish urine observed between week 2 and sacrifice. Increased prothrom bin times in males and increased activated partial throm boplastin times in both sexes.
Intermediate- Term Inhalation (1 to 6 m onths)	O ral study NOAEL=30 mg/kg/day (inhalation absorption rate= 100%)	OccupationalLOC forMOE = 100	Subchronic feeding study in the rat Reddish urine observed between week 2 and sacrifice. Increased prothrom bin times in males and increased activated partial throm boplastin times in both sexes.

Hum an Exposures and Risks

- A. Residential: No residential exposure is expected, since the pesticide is not foruse in or around residential areas.
- B. CommercialHandlers: Data from the Pesticide Handler Exposure Database (PHED) were used to estimate exposure of handlers to acequinocyl.

Short T erm: The combined M argin of Exposure $(M \circ E)$ for the most highly exposed handler in an ornam ental greenhouse setting (a single individual whom ixes, bads, and applies the materials using high-pressure, hand-wand equipment) is 3,397, which is well over the target $M \circ E$ of 100. Based primarily upon the proposed use practices, the Agency expects that,

typically, com m ercial and private (i.e., grow er) pesticide handlers will experience short-term exposures (1 - 30 days). The label directs that sequential applications should not be made.

Interm ediate: A lithough the Agency does not typically expect interm ediate-term handler exposures, the risks estim ated for short-term exposures are conservative and adequate to protect handlers who m ight experience interm ediate-term exposures. As stated above under Short-Term, the estimated exposure and risk to a mixer/bader/applicator using high pressure, hand-wand equipment on greenhouse ornamentals are well over the target MOE of 100.

C. Post-Application; Agricultural Workers:

Short Term: The transfer coefficient data from A gricultural Re-Entry Task Force (ARTF) studies were used in conjunction with the applicant-submitted chemical-specific disbolgeable foliar residue (DPR) dissipation data from chrysanthem um s grown in a greenhouse. The DPR study evaluated acequinocyland its metabolite, acequinocyl-OH (deacetylated acequinocyl). The resulting MOE was greater than 33,000, corrected to reflect the 2X maximum label rate used in the DFR study. The Agency expects post-application agricultural exposures to workers would typically be short-term (1-30 days). Mites are typically "hotspot" (i.e., relatively small areas) pests; therefore, entire facilities typically need not be treated form ites. A cequinocyl is not to be applied sequentially.

Interm ediate Term: A lithough the Agency does not typically expect interm ediate-term handler exposures, the risks estimated for short-term exposures are conservative and adequate to protect workers who might experience in term ediate-term exposures. The short-and intermediate-term dermal and inhalation endpoints are the same, and do not exceed EPA's level of concern.

Environm entalCharacteristics

The environmental fate data supportive of the non-food use of acequinocyl in greenhouses and shadehouses are described below:

- A . Hydrolysis: Based on the submitted supplemental study, acequinocyl undergoes rapid hydrolysis underneutral and alkaline pH conditions and is considered stable underacidic conditions. Deacetylated acequinocyl was reported to be the majorm etabolite.
- B. A erobic Soil M etabolism: Based on the available study, observed half-lives of degradation in the sandy barn and silt soils were less than 2 days. Deacetylated acequinocyl was reported to be the major metabolite.
- C. Adsorption/Desorption/Leaching: Based on the available studies, acequinocylis expected to exhibit low mobility in soiland is expected to exhibit low potential to leach to ground water.

Ecological Characteristics/Risk

The ecological effects data considered in the qualitative risk assessment for the proposed use in greenhouses/shadehouses are described below. A quantitative ecological risk assessment of the proposed greenhouse use is not needed, since limited environmental exposure is expected from this indooruse.

- A. Birds: It is not expected that the proposed use of acequinocyl would result in exposure levels that would be of concern to birds. Based on the available acute avian studies, acequinocyl is characterized as practically non-toxic to moderately toxic to birds.
- B. Fish: It is not expected that adverse effects to fish would result from the proposed use of acequinocyl, due to its use indoors and the fact that acequinocyl's solubility lim it is 2-3 orders of magnitude lower than the concentration at which the toxic effects occur. Additional acute toxicity testing is being requested, using the metabolite itself to ascertain whether the reported toxic effects are due to parent, metabolite, or both. Based on the results of the available acute fish studies, acequinocyl is characterized as slightly (formulated product tested) to moderately (technical product tested) to freshwater fish, and slightly (formulated product tested) to highly toxic (technical product tested) to estuarine/marine fish.
- C. A quatric Invertebrates: Further acute toxicity testing for aquatric invertebrates is needed to fully characterize the toxicity to invertebrates. The additional testing includes (1) testing on either daphnids or mysids using deacety lated acequinocyl itself to ascertain whether the toxic effects reported are due to the parent, the deacety lated acequinocyl, or both, and (2) testing with the metabolite in a sediment toxicity test using freshwater invertebrates as the test organism. Results of the available acute freshwater invertebrate study conducted with the formulated product show acequinocyl to be slightly toxic, and a study using the technical product showed very high toxicity to the test species. In the available estuarine/invertebrate studies, both conducted with the technical product, acequinocyl was shown to be very highly toxic to the test species.

A dditional studies are to be conducted on the end-use form ulation to further characterize its toxicity to aquatic invertebrates. To m itigate the risk to aquatic invertebrates during the period that the studies are in progress, labeling will be required in connection with the Environmental Hazards to advise the user that the pesticide is very highly toxic to invertebrates, such as Eastern oysters and mysid shrimp, and that drift and runoff may be hazardous to aquatic organisms in water adjacent to treated areas. In addition, the end-use product is not expected to present a significant risk to invertebrates, based on its lower toxicity (slightly toxic) and the nature of the use (indoors in commercial greenhouses/shadehouses).

D. M am m als: It is not expected that the proposed use of acequinocyl would result in exposure levels that would be of concern to m am m als. M am m alian studies show acequinocyl to be of very low toxicity to m am m als.

4. <u>SUM M ARY OF REGULATORY POSITION AND RATIONALE</u>

Based on the available data as described in this docum ent, there is adequate inform ation to support a registration decision under FIFRA section 3 (c) (7) (C) for the conditional registration of the pesticide

products, acequinocyl technical and the form ulated product described in this document, for use on ornam ental plants grown in comm encial greenhouses and shadehouses.

Use, Form ulation, Manufacturing Process or Geographic Restrictions

The form ulated product falls under the scope of the W orker Protection Standard (W PS) and must be used only in accordance with its labeling and the W PS, 40 CFR part 170. The form ulated product label includes the W PS restrictions cited below.

A restricted entry interval of 12 hours.

The standard W PS drift restriction of not applying in any way that will contact workers or other persons, either directly or through drift, and the statem entithat only protected workers may be in the area during application.

Use of Personal Protective Equipm ent (PPE) by applicators and other handlers, consisting of long-sleeved shirt and long pants, socks, shoes and chem ical resistant gloves made of waterproof material.

The label bears the following additional use restriction statem ents.

Not foruse in orazound residential sites. Foruse only in commercial greenhouses and shadehouses.

Restrictions for Use on O mam ental Crops Grown in Greenhouses:

- < Do not apply through any type of irrigation system.
- < Do not contam inate water, food or feed by storage or disposal.
- < DO NOT contam inate waterwhen disposing of equipment washwaters.

5. SUM M ARY OF DATA GAPS

The additional data listed below must be provided to the Agency as conditions of registration. Upon receipt and evaluation of the requested information, the Agency will reassess the registration, and, if appropriate, will remove these conditions.

- < One year storage stability and comosion characteristics (technical)
- < Skin sensitization study (end-use product)
- < Additional data to upgrade the aerobic soil metabolism study
- < Additional data to upgrade the hydrolysis study
- < A dsorption/desorption study

- < Freshwater invertebrate (Daphnia) study
- < A cute sedim ent toxicity test for freshwater invertebrates conducted with the majorm etabolite
- < A cute toxicity to freshwater fish study conducted with the majorm etabolite
- < A cute toxicity to aquatic invertebrates study conducted with the majorm etabolite

6. <u>CONTACT PERSON AT EPA</u>

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